Title: RNA Aptamers and Methods for Identifying the Same Applicant(s): Sullenger et al.

Attorney Docket No.: 180/124/2

FIGURE 1A

- 9.1 (SEQ ID NO:1): 5' GGGAGAGAG AAGAGGGAUG GG CCGCCAGU GGGAAGCUAU ACCCAACGCC CCAGCCCCAG AGCAUAACCC AGAGGUCGAU AGUACUGGAU CCCCCC 3'
- 9.2 (SEQ ID NO:2): 5' GGGAGAGAG AAGAGGGAUG GGCUAUAUAC ACGCUGGUGA UCCCAUCUCA AUUGAAACAA CACAUAACCC AGAGGUCGAU AGUACUGGAU CCCCCC 3'
- 9.3 (SEQ ID NO:3): 5' GGGAGAGAGG AAGAGGGAUG GGGACUAUAC CGCGUAAUGC UGCCUCCCCA UUCCGGAACG CUCAUAACCC AGAGGUCGAU AGUACUGGAU CCCCCC 3'
- 9.4 (SEQ ID NO:4): 5' GGGAGAGAG AAGAGGGAUG GGCACUAUAC GCAUCUUGCU GCCUGCCCGC GAGUCAAAUU GCAUAACCCA GAGGUCGAUA GUACUGGAUC CCCCC 3'
- 9.5 (SEQ ID NO:5): 5' GGGAGAGAG AAGAGGGAUG GGCCUACCAG UUCGUGGCUA GCGUGACGUA CCACCCAGGG ACCAUAACCC AGAGGUCGAU AGUACUGGAU CCCCCC 3'
- 9.7(SEQ ID NO:6): 5' GGGAGAGAG AAGAGGGAUG GGCGAUAACC AACAUGGUGA UCCCAUUCAU CAUACCCUAC AACAUAACCC AGAGGUCGAU AGUACUGGAU CCCCCC 3'
- 9.8 (SEQ ID NO:7): 5' GGGAGAGAG AAGAGGGAUG GGGCCACCUA CUAUACCGGU CAUCGUGCAU AGGUCGCUGC CACAUAACCC AGAGGUCGAU AGUACUGGAU CCCCCC 3'
- 9.9(SEQ ID NO:8): 5' GGGAGAGAGG AAGAGGGAUG GGUCUCACAC CCGAAGAUGG CCAAAGAGGG AGAUGAGUUU CCAUAACCCA GAGGUCGAUA GUACUGGAUC CCCCC 3'
- 9.11(SEQ ID NO:9): 5' GGGAGAGAGG AAGAGGGAUG GGACUAUAUU CGGAAUCUGG ACUCCCACCU GCCUGCCCCA GACAUAACCC AGAGGUCGAU AGUACUGGAU CCCCCC 3'
- 9.12(SEQ ID NO:10): 5' GGGAGAGAG AAGAGGGAUG GGCGAUAUAC ACAUUGGUGA UCCCACCCAC AUGAAACCAC AGCAUAACCC AGAGGUCGAU AGUACUGGAU CCCCCC 3'
- 9.13 (SEQ ID NO:11): 5' GGGAGAGAG AAGAGGGAUG GGCUCAUCAC AGGCGAAGUG AACAACACUA CCGNCNAGUU ACCAUAACCC AGAGGUCGAU AGUACUGGAU CCCCCC 3'
- 9.14 (SEQ ID NO:12): 5' GGGAGAGAGG AAGAGGGAUG GG GACUAUAC GUGAACGACU GCAUCCACUUC CCGCCAUGG CAUAACCCAG AGGUCGAUAG UACUGGAUCC CCCC 3'

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FIGURE 1B

- 9.16(SEQ ID NO:13): 5' GGGAGAGAG AAGAGGGAUG GGCCAUACGU GGACGACUGC ACCCGACCCU UCAGCCCAGG UCCAUAACCC AGAGGUCGAU AGUACUGGAU CCCCCC 3'
- 9.17(SEQ ID NO:14): 5' GGGAGAGAG AAGAGGGAUG GGACCAUACG CACAUUGCUG AAUCCCCCUC AAUAGCACCU ACCAUAACCC AGAGGUCGAU AGUACUGGAU CCCCCC 3'
- 9.18 (SEQ ID NO:15): 5' GGGAGAGAG AAGAGGGAUG GGCCAUAACC ACUUUGGUGA ACCCACCCAG CUCc/UUGUGAU UGCAUAACCC AGAGGUCGAU AGUACUGGAU CCCCC 3'
- 9.19(SEQ ID NO:16): 5' GGGAGAGAG AAGAGGGAUG GGACCAUAAC GACUACUCGUGA AUCCCACCAU CAGCGCACAA CAUAACCCAGA GGUCGAUAG UACUGGAUCC CCCC 3'
- 9.20(SEQ ID NO:17): 5' GGGAGAGAG AAGAGGGAUG GGGACUAUAC CGGCÄAUCGU GCAUCCCUG GACCUAACAA UACAUAACCC AGAGGUCGAU AGUACUGGAU CCCCCC 3'
- 9.21(SEQ ID NO:18): 5' GGGAGAGAGG AAGAGGGAUG GG AACACCAU UAAUGCUCGG CCAGGUAACC CCGGCGCAUA CUCAUAACCC AGAGGUCGAU AGUACUGGAU CCCCC 3'
- 9.25(SEQ ID NO:19): 5' GGGAGAGAG AAGAGGGAUG GGGACCAUAA CUCUAACGGG UGAAUCCCGC AUCUCGACAA UACAUAACCC AGAGGUCGAU AGUACUGGAU CCCCCC 3'
- 9.26(SEQ ID NO:20): 5' GGGAGAGAGG AAGAGGGAUG GG UGAUAACC ACUCUGGUGA ACCCCUCCCG ACUUGCUCGC ACAUAACCCA GAGGUCGAUA GUACUGGAUC CCCCC 3'
- 9.27(SEQ ID NO:21): 5' GGGAGAGAGG AAGAGGGAUG GGUAAUAACU GUAUGGUGAA CCCACCCAAA CUCCCAUGGC UACAUAACCC AGAGGUCGAU AGUACUGGAU CCCCCC 3'
- 9.28 (SEQ ID NO:22): 5' GGGAGAGAG AAGAGGGAUG GG CGCCAUAC GCACAUUGCU GCAUCGCCUU CCCGUAAGAA CCAUAACCCA GAGGUCGAUA GUACUGGAUC CCCCC 3'

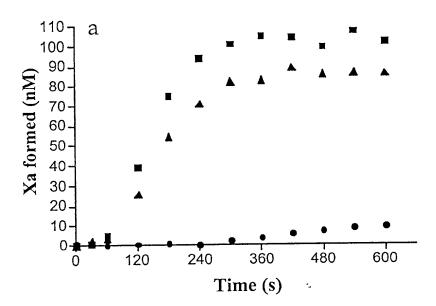
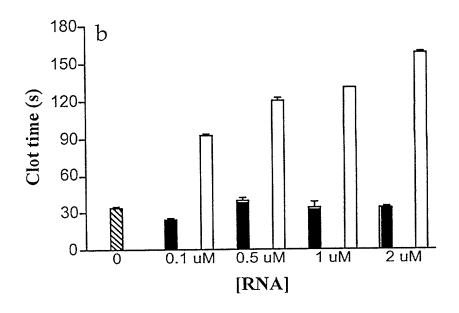


FIGURE 2A

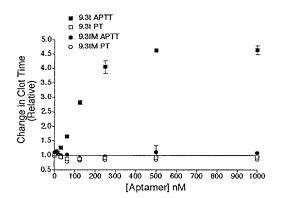
FIGURE 2B



RNA Aptamers and Methods for Identifying the Same t(s): Sullenger et al. Title:

Applicant(s): Attorney Docket No.:

180/124/2



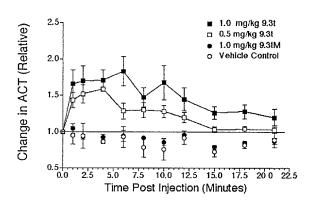


Figure 3A

Figure 3B

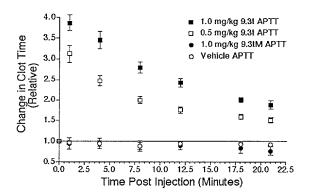
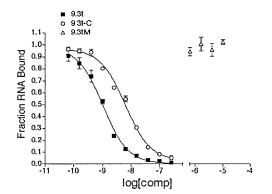


Figure 3C

RNA Aptamers and Methods for Identifying the Same It(s): Sullenger et al.
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Applicant(s): Sul Attorney Docket No.:



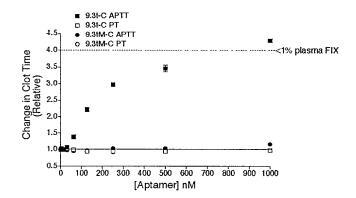


Figure 4A

Figure 4B

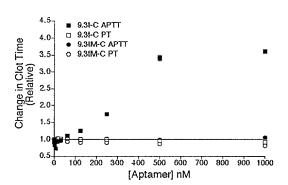


Figure 4C

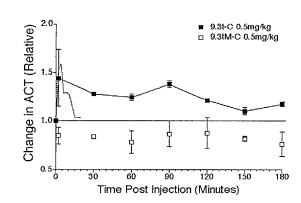


Figure 5A

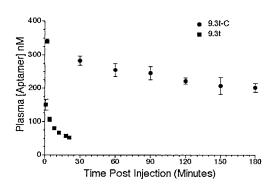


Figure 5B

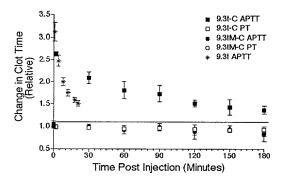


Figure 5C

Title: RNA Aptamers and Methods for Identifying the Same

Applicant(s): Sullenger et al. Attorney Docket No.: 180/124/2

Figure 6A

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- 10.1(SEQ ID NO:23): 5' GGGAGAGGG AAGAGGGAUG GGAAAAUAGC CCCAGCGAGA UAAUACUUGG CCCCCGUACCA CCAUAACCCA GAGGUCGAUA GUACUGGAUC CCCCC 3'
- 10.5(SEQ ID NO:24): 5' GGGAGAGG AAGAGGGAUG GGCCAGAAGG AACUAAACAC CUGAACCCC CAUCGCGAGAG ACCAUAACCC AGAGGUCGAU AGUACUGGAU CCCCCC 3'
- 10.6(SEQ ID NO:25): 5' GGGAGAGAG AAGAGGGAUG GGAUGUCACU UGGCCCCUCG CGCACC/aCGCC AGCGAGCCCA UAACCCAGAG GUCGAUAGUA CUGGAUCCCC CC 3'
- 10.7(SEQ ID NO:26): 5' GGGAGAGAGG AAGAGGGAUG GGACACGCCC AGCGAGCUCA AACUUGGCCC CCGUGCAUCA CC CCAUAACC CAGAGGUCGA UAGUACUGGA UCCCCCC 3'
- 10.8(SEQ ID NO:27): 5' GGGAGAGGG AAGAGGGAUG GGAAGUGCCA CAGCGAGCAC AUGACUUGGC CCCGCAUUGC ACCCAUAACC CAGAGGUCGA UAGUACUGGA UCCCCCC 3'
- 10.11(SEQ ID NO:28): 5' GGGAGAGGG AAGAGGGAUG GGAAACUAAU GCCCUAGCGA GCAUACCCGG ACUGGCCCCG CCAUAACCC AGAGGUCGAU AGUACUGGAU CCCCCC 3'
- 10.12(SEQ ID NO:29): 5' GGGAGAGAG AAGAGGGAUG GGAAAAUAGC
 CCCAGCGAGA UAAUACUUGG CCCCGCUACU ACCCAUAACC CAGAGGUCGA UAGUACUGGA
 UCCCCCC 3'
 - 10.13(SEQ ID NO:30): 5' GGGAGAGGG AAGAGGGAUG GGCGACCCCA CUGGCGGAAA CCGACAAUCA CUCCCCACGA CCAUAACCC AGAGGUCGAU AGUACUGGAU CCCCCC 3'
 - 10.14 (SEQ ID NO:73): 5'GGGAGAGGG AAGAGGGAUG GGAAAAUAGC CCCAGCGAGA UAAUACUUGG CCCCGCUACU ACCAUAACCC AGAGGUCGAU AGUACUGGAU CC 3'

Title: RNA Aptamers and Methods for Identifying the Same

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Figure 6B

- 10.15(SEQ ID NO:31): 5' GGGAGAGAG AAGAGGGAUG GGCAGCCCAG CGAGGGACAC UUAACCCCCU GUCCCCCAUC CAAACCAUAA CCCAGAGGUC GAUAGUACUG GAUCCCCCC 3'
- 10.18(SEQ ID NO:32): 5' GGGAGAGGG AAGAGGGAUG GGCCAGAAGU CACCGCGACG GUACUGAACC CCCCACCCAA ACCCAUAACC CAGAGGUCGA UAGUACUGGA UCCCCCC 3'
- 10.19(SEQ ID NO:33): 5' GGGAGAGAG AAGAGGGAUG GGCCAGAAGU GCUCACUACA ACGCUUUGAC CCCCCCAUCC ACAUCCCAUA ACCCAGAGGU CGAUAGUACU GGAUCCCCCC 3'
- 10.21(SEQ ID NO:34): 5' GGGAGAGGG AAGAGGGAUG GG CCAGCAAC CGAAGGGCGG AAUACCCCCC GUCUCCACAU ACCCAUAACC CAGAGGUCGA UAGUACUGGA UCCCCCC 3'
- 10.22(SEQ ID NO:35): 5' GGGAGAGAG AAGAGGGAUG GG ACGCGACU CAGGCAGCAC UUGACUUGGC CCCUUGCGAU CACCAUAACC CAGAGGUCGA UAGUACUGGA UCCCCC 3'
- 10.23 (SEQ ID NO:36): 5' GGGAGAGGG AAGAGGGAUG GG CCAGCAAC GCUAACACGG AAUACCCCCC ACCCCAACGU GCCCAUAACC CAGAGGUCGA UAGUACUGGA UCCCCCC 3'
- 10.24(SEQ ID NO:37): 5' GGGAGAGAG AAGAGGGAUG GG CUUCUCAA CCGAAAUACA ACUUUAAAUC AUUUAUCACU UACCAUAACC CAGAGGUCGA UAGUACUGGA UCCCCC 3'
- 10.30(SEQ ID NO:38): 5' GGGAGAGAGG AAGAGGGAUG GGAUACGCCG AUGCAAGCAU GUCCACACAC CGCAUGCCGU ACCCAUAACC CAGAGGUCGA UAGUACUGGA UCCCCCC 3'

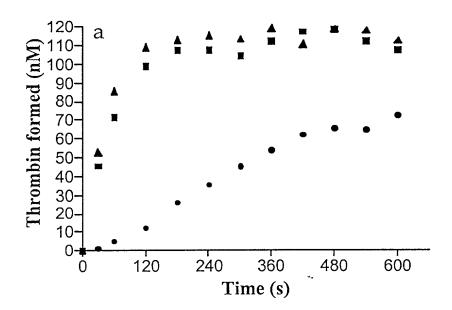
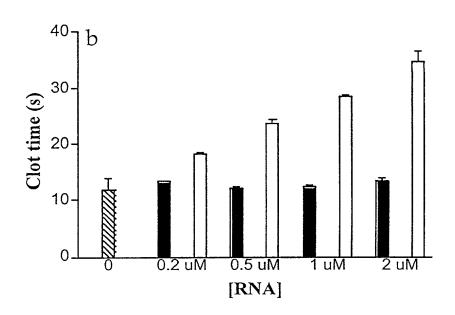


Figure 7A Figure 7B



RNA Aptamers and Methods for Identifying the Same

Applicant(s): Sullenger et al. Attorney Docket No.:

180/124/2

RNA:Protein Complexes

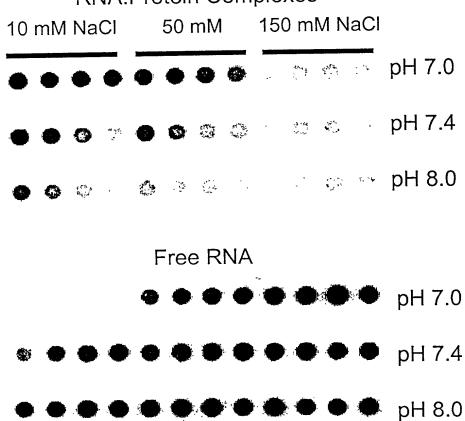
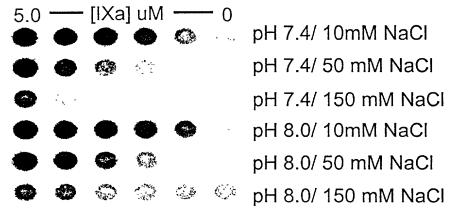


FIGURE 8

Title: RNA Aptamers and Methods for Identifying the Same Applicant(s): Sullenger et al.

Attorney Docket No.: 180/124

RNA:Protein Complexes



Free RNA

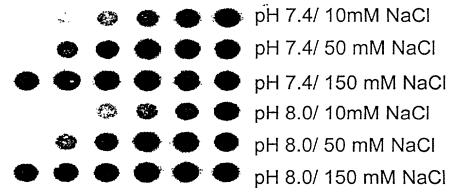


FIGURE 9

Title: RNA Aptamers and Methods for Identifying the Same

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Figure 10

16.1: (SEQ ID NO:39) 5' GGGAGAGAGG AAGAGGGAUG GGUACAGAGG AGUACAAGUA GCAUGGUCCC CUCGUGUAAA AACAUAACCC AGAGGUCGAU AGUACUGGAU CCCCCC 3'

16.2 (SEQ ID NO:40): 5' GGGAGAGAGG AAGAGGGAUG GGUGCAAAAG AGCUUCUUGU AGUAUGAUCC CUCAACCGCA AGCAUAACCC AGAGGUCGAU AGUACUGGAU CCCCCC 3'

16.3 (SEQ ID NO:41): 5' GGGAGAGAGG AAGAGGGAUG GG UACAGAGG AGUACAAGUA GCAUGAUCCC CUCGUGUAAA AACAUAACCC AGAGGUCGAU AGUACUGGAU CCCCCC 3'

16.5 (SEQ ID NO:42): 5' GGGAGAGAGG AAGAGGGAUG GGAGCCUAUG UAACAGAUGC AGAUCCCUAG UCGUCCCAAC ACCAUAACCC AGAGGUCGAU AGUACUGGAU CCCCCC 3'

16.7 (SEQ ID NO:43): 5' GGGAGAGAGG AAGAGGGAUG GGCACAACGA ACACCGCAUC CCUUGACAGA AAGAGCACGC CUCAUAACCC AGAGGUCGAU AGUACUGGAU CCCCCC 3'

16.10 (SEQ ID NO:44): 5' GGGAGAGAGG AAGAGGGAUG GGUACAGAGG AGUACAAGUA ACAUGAUCCC CUCGUGUAAA AACAUAACCC AGAGGUCGAU AGUACUGGAU CCCCCC 3'

16.11 (SEQ ID NO:45): 5' GGGAGAGAGG AAGAGGGAUG GG CACAACGA ACACCGCAUC CCUUGACAGA AAGAACACGC CUCAUAACCC AGAGGUCGAU AGUACUGGAU CCCCCC 3'

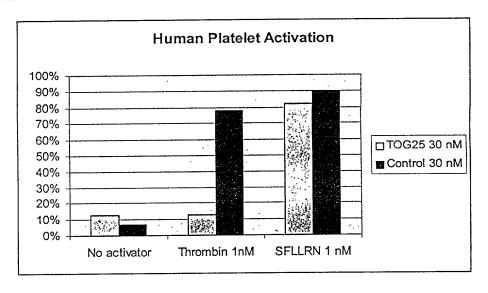
16.18 (SEQ ID NO:46): 5' GGGAGAGAGG AAGAGGGAUG GGCACAAGGA ACACCGCAUC CCUUGACAGA AAGAACACGC CUCAUAACCC AGAGGUCGAU AGUACUGGAU CCCCCC 3'

16.20 (SEQ ID NO:47): 5' GGGAGAGAGG AAGAGGGAUG GGAGCCUAUG UAACAGAUGC AGAUCCCUAG ACGACCCAAC ACCAUAACCC AGAGGUCGAU AGUACUGGAU CCCCCC 3'

FIGURE 11

Random region sequences (Round 13)	K _d Human Thrombin	K _d Porcine Thrombin
PIG 5 UGCGAACAAAGCUGAAGUACUUACGCACAACCCGUAGAAU	3 nM	1 nM
PIG 7 AACAACUGAAGAACUACCCUUCUUACUGACGAAUUAA	1 nM	<0.5 nM
PIG 8 AAACAAAGCUGAAGUACUUAUUCCAUCACCACGCCGGAA	l nM	0.5 nM
PIG 10 UAUUUGGCUUCUCAGUGCCGCAGAGACAGCAACAAUUAGU	>>50 nM	0.5 nM
HUMAN ACAAAGCUGGAGAACUUACCGUUCCCUCUCCAGAGAUCAA	2 nM	0.5 nM
TOGGLE 25 GAACAAAGCUGAAGUACUUACCCAAGAUCAUCCCGAACGA	5 nM	0.5 nM
TOGGLE 30 AACAAAGCUGGAGAACUUAACGUCCCUCUCCCAGCGGUAA	3 nM	0.5 nM

FIGURE 12



Title: RNA Aptamers and Methods for Identifying the Same

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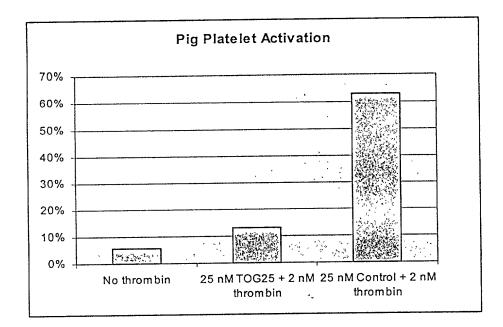


FIGURE 13

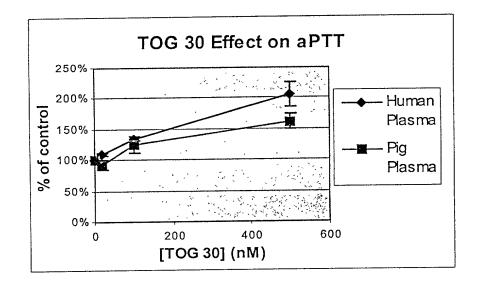


FIGURE 14

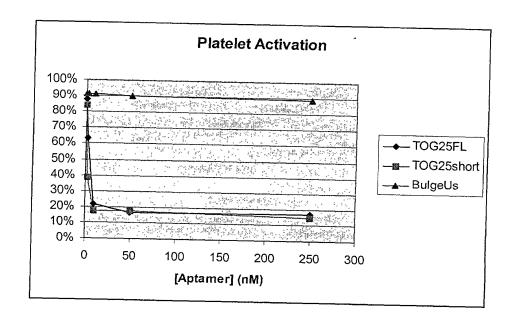


| Loop U2 | Kd ~ 0.5 nM NoStem2 Ka>> 1000 nM NoStem1 Ka~ 1000 nM 8 Q \Box Ω

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Figure 16



Title: RNA Aptamers and Methods for Identifying the Same Applicant(s): Sullenger et al.

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Figure 17

E2F-3 Round 10 Aptamers

5' Primer GGG AGA GAG GAA GAG GGA NGG G (SEQ ID NO: 62)

3' Primer CAUA ACC CAG AGG UCG A A GUA CUG GAUCCC CCC (SEQ ID NO: 63)

10-1 and 10-8 (SEQ ID NO: 64)
5' P-AAUGGA AUC ACUGAA GGC CCUCCG UAG CAC CUA ACA CAG U-3' P

10-2 (SEQ.ID NO: 65)
5' P-GCAVICC VIGC CAG CGG CGA CGG ACC VIVC GCC CAC AGG CC'VC-3' P

10-3, 10-7, 10-11, and 10-12 (SEQ ID NO: 66)
5' P-UUA 'UA AGC ACA CUG AAG CCC UCA GCA AAA CCU CCA CAG G-3' P

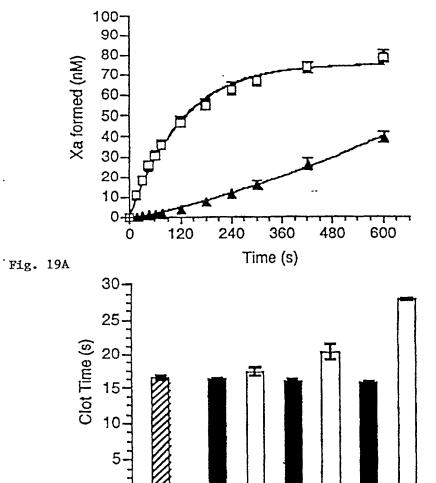
10-4 (SEQ ID NO: 67)
5' P-UAU GAA AUC ACA GAA GCC CGC GUU CGA CAC CUC CAC UGUU 3' P

10-5 (SEQ ID NO: 68)
5' P-CAA AC LICAC AGA CUE CAA CUE CAG GAG CAC CCA CCC ACA CUE
GGA CAG-3' P

10-6 (SEQ ID NO: 69)
5' P-AUC CCC GCC GUA AGC CGU CCU GAU GGA CAC CAC ACU CCG C-3' P

	S1	L1	S2	L2	S2	L3	S1
*9-3 5' *9-20 5' *9-25 5' *9-26 5' *9-28 5' *9-17 5' 9-17 5' 9-18 5' 9-18 5' 9-18 5' 9-18 5' 9-18 5' 9-18 5' 9-14 5' 9-14 5' 9-17 5'	gggaugggGA augggGA gggaugggGA gggaugggCG gaggaugggA gaggaugggA gaggaugggA gggaugggCA aggaugggCA aggauggg ggauggg ggauggg ggauggg ggauggg ggauggg ggauggg ggauggg gggauggg	CUAUACC CUAUACCG CCAUA UGAUA CCAUAC CCAUA CCAUAC CUAUA CUAUAC CCAUA CUAUA CUAUA CUAUA CUAUAUA CUAUAUA CUAUAUA	GCG GCA ACUC ACCA GCA ACGA GCA UUCGG GCA CGU ACCA ACCA	UAAUGC AUCG UAAC CUC CAU CUAC CAU AAU UCU GG CUU ACA ACAUUG GCUG GAACG GUA	UGC GGGU UGC UCGU UGC CUGGA UGC ACG UGGU UGGU	C A GAA GAA UGCAU GAA UGAA ACUGCA GAA GAU AU AU GCA UGAA	UCCCAUUCC UCCCCU UCCCCUCC CGCCUUCCC UCCCACCAUC UCCCACCAUC UCCCACCU CUCCACCU CCCACCC
The fluid time time the fluid time the fluid time the fluid time time time time time time time time			FIGURE	18			

FIGURE 18



125 nM

0

2.5 μΜ

[RNA]

10 μM

Fig. 19B

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Figure 20

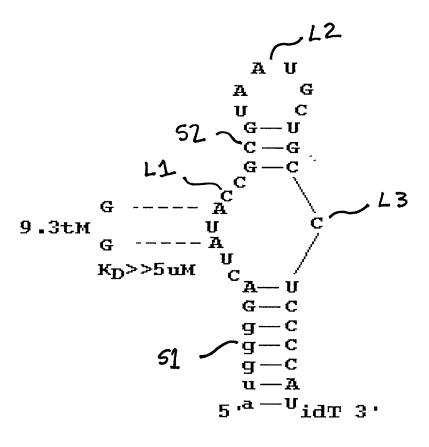
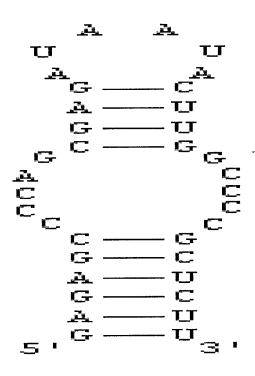


Figure 21



RNA Aptamers and Methods for Identifying the Same tt(s): Sullenger et al.

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Applicant(s): Sul Attorney Docket No.:

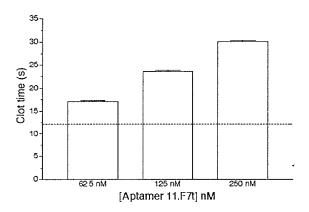


Figure 22A

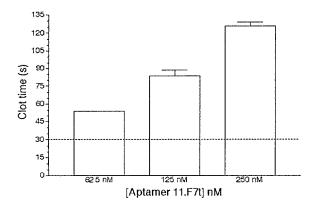


Figure 22B

Figure 23A

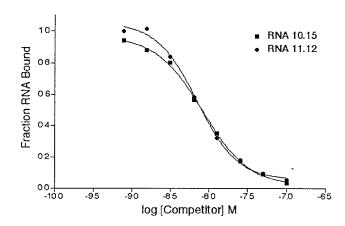
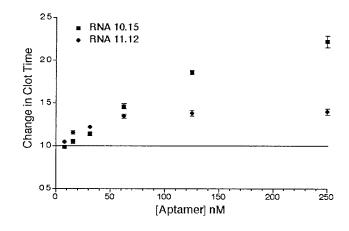


Figure 23B



itle: RNA Aptamers and Methods for Identifying the Same

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Figure 24

ANG9-4 Binding

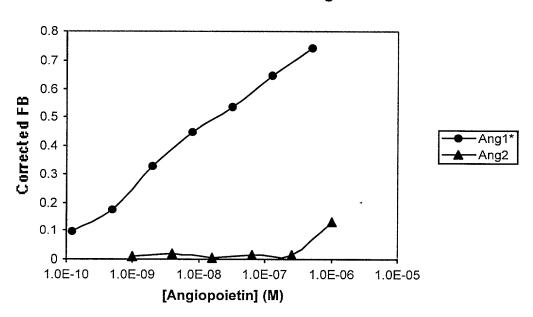
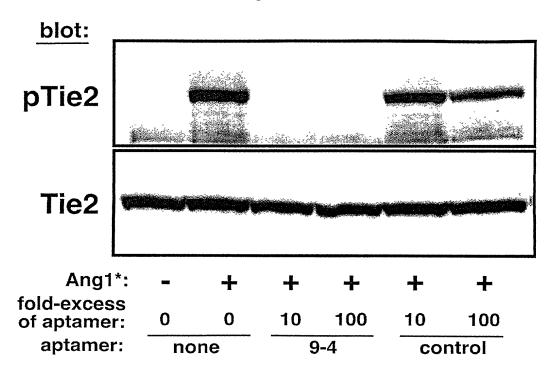


Figure 25



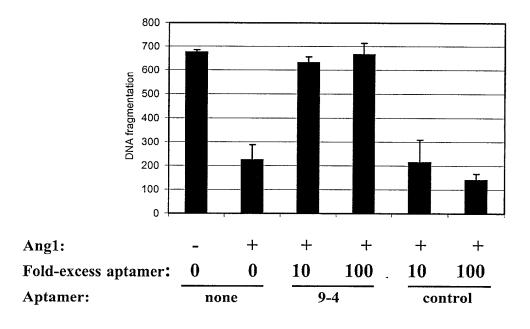


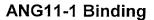
Figure 26

The first term that the first term of the first

Ang2:

Aptamer:

Sullenger et al. No.: 180/124/2 Applicant(s): Attorney Docket No .:



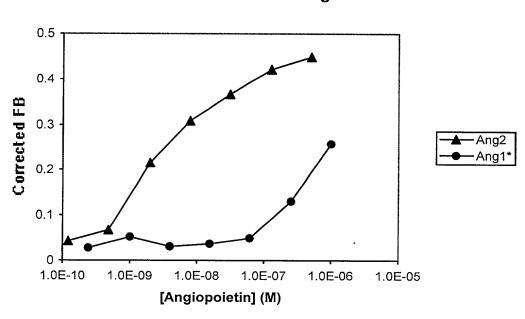


Figure 27

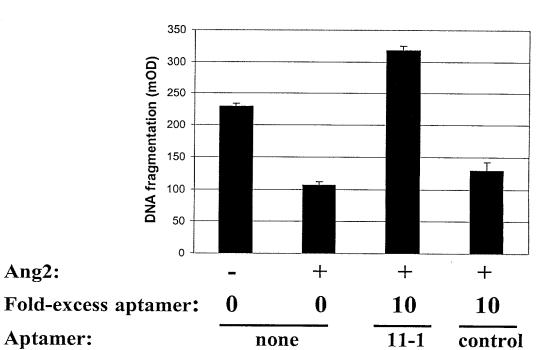


Figure 28

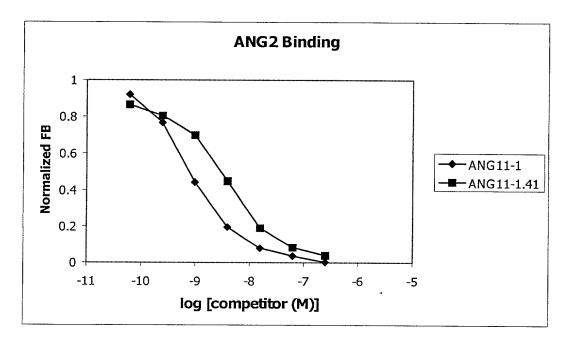


Figure 29

plt22jpg by D. Stewart and M. Zuker C 2001 Washington University

dG = -1.43 (initially -4.2] ANG11-1.41

Figure 30

Figure 31

